

WHAT IS CLAIMED IS:

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1. An isolated nucleic acid encoding an osteoprotegerin binding protein selected from the group consisting of:
 - a) the nucleic acid sequence as in Figure 1 (SEQ ID NO:____) and Figure 4 (SEQ ID NO:____);
 - b) nucleic acids which hybridize to the polypeptide coding regions as shown in Figure 1 (SEQ ID NO:____) and Figure 4 (SEQ ID NO:____) and remain hybridized under high stringency conditions; and
 - c) nucleic acids which are degenerate to the nucleic acids of (a) or (b).
 - 15 2. The nucleic acid of Claim 1 which is cDNA, genomic DNA, synthetic DNA or RNA.
 - 20 3. A polypeptide encoded by the nucleic acid of Claim 1.
 - 25 4. The nucleic acid of Claim 1 including one or more codons preferred for Escherichia coli expression.
 - 30 5. The nucleic acid of Claim 1 having a detectable label attached thereto.
 - 35 6. A nucleic acid encoding a polypeptide comprising the amino acid sequence of residues 1-316 and residues 70-316 as shown in Figure 1 (SEQ ID NO: ____).
 7. A nucleic acid encoding a polypeptide comprising amino acid sequence of residues 1-317 and residues 69-317 as shown in Figure 4 (SEQ ID NO: ____);

8. A nucleic acid encoding a soluble osteoprotegerin binding protein.

9. The nucleic acid of Claim 8 encoding a polypeptide comprising residues 69-317 as shown in Figure 4 (SEQ ID NO:____) and truncations thereof;

10. An expression vector comprising the nucleic acid of Claims 1 and 9.

11. The expression vector of Claim 10 wherein the nucleic acid comprises the polypeptide-encoding region as shown in Figure 1 (SEQ ID NO:____) and Figure 4 (SEQ ID NO:____);

15. 12. A host cell transformed or transfected with the expression vector of Claim 10.

20. 13. The host cell of Claim 12 which is a eucaryotic or procaryotic cell.

14. The host cell of Claim 13 which is Escherichia coli.

25. 15. A process for the production of an osteoprotegerin binding protein comprising:
growing under suitable nutrient conditions host cells transformed or transfected with the nucleic acid of Claim 1; and
30. isolating the polypeptide product of the expression of the nucleic acid.

35. 16. A polypeptide produced by the process of Claim 15.

17. A purified and isolated osteoprotegerin binding protein, or fragment, analog, or derivative thereof.

5 18. The protein of Claim 17 which is a human osteoprotegerin.

10 19. The protein of Claim 17 having the amino acid sequence as shown in Figure 1 (SEQ ID NO:____) and Figure 4 (SEQ ID NO:____).

20. The protein of Claim 17 which has been covalently modified with a water-soluble polymer.

15 21. The protein of Claim 20 wherein the polymer is polyethylene glycol.

20 22. The protein of Claim 17 which is a soluble osteoprotegerin binding protein.

25 23. The protein of Claim 22 comprising the amino acid sequence from residues 70-316 inclusive as shown in Figure 1 (SEQ ID NO: ____), or a fragment, analog, or derivative thereof.

25 24. The protein of Claim 22 comprising the amino acid sequence from residues 69-317 inclusive as shown in Figure 4 (SEQ ID NO: ____) and truncations thereof.

30 25. An antibody or fragment thereof which specifically binds an osteoprotegerin binding protein.

35 26. The antibody of Claim 25 which is a monoclonal antibody.

27. A method for detecting the presence of an osteoprotegerin binding protein in a biological sample comprising:

5 incubating the sample with the antibody of Claim 25 under conditions that allow binding of the antibody to the osteoprotegerin binding protein; and detecting the bound antibody.

10 28. A method for detecting the presence of osteoprotegerin in a biological sample comprising:

incubating the sample with an osteoprotegerin binding protein under conditions that allow binding of the protein to osteoprotegerin; and

15 measuring the bound osteoprotegerin binding protein.

29. A method to assess the ability of a candidate compound to bind to an osteoprotegerin binding 20 protein comprising:

incubating the osteoprotegerin binding protein with the candidate compound under conditions that allow binding; and

measuring the bound compound.

25 30. The method of Claim 29 wherein the compound is an agonist or an antagonist of an osteoprotegerin binding protein.

30 31. A method of regulating expression of an osteoprotegerin binding protein in an animal comprising administering to the animal a nucleic acid complementary to the nucleic acids as shown in Figure 1 (SEQ ID NO: __) and Figure 4 (SEQ ID NO: __).
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32. A pharmaceutical composition comprising a therapeutically effective amount of an osteoprotegerin binding protein in a pharmaceutically acceptable carrier, adjuvant, solubilizer, stabilizer and/or 5 anti-oxidant.

33. The composition of Claim 32 wherein the osteoprotegerin binding protein is a human osteoprotegerin binding protein.

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34. A method of treating bone disease in a mammal comprising administering a therapeutically effective amount of a modulator of an osteoprotegerin binding protein.

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35. The method of Claim 34 wherein the modulator is a soluble form of an osteoprotegerin binding protein.

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36. The method of Claim 35 wherein the modulator is an antibody, or fragment thereof, which specifically binds an osteoprotegerin binding protein.

Add A.7

Add C5

Add E1